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Subject: BAVAP#09042488

Ready to allow:

A method for modulating the expression of an exogenous gene in an isolated cell containing:

(i) a modified ecdysone receptor which, in the presence of a ligand therefor, and optionally in the further presence of a silent partner therefor, binds to a response element wherein said modified ecdysone receptor comprises:

- (a) a ligand binding domain that binds to an ecdystroid,
- (b) a DNA-binding domain obtained from a DNA-binding protein, which binds to said response element; and
- (c) an activation domain of a transcription factor,

wherein at least one of said DNA-binding domain or said activation domain is not obtained from a native ecdysone receptor, with the proviso that when said activation domain is derived from a glucocorticoid receptor, said DNA-binding domain is not derived from a glucocorticoid receptor or an E. ooll LexA protein; and

(ii) a DNA construct comprising said exogenous gene under the control of said response element, wherein said response element:

- (a) is a modified response element which comprises, a first half-site and a second half-site separated by a spacer of 0-5 nucleotides; and wherein said second half-site is obtained from a glucocorticoid receptor subfamily response element
- (b) binds to said modified ecdysone receptor, and
- (c) does not bind to Farnesoid-X receptor (FXR);

said method comprising providing to the cell an effective amount of one or more ligands for said modified ecdysone receptor; wherein said one or more ligands are not normally present in the cell; and wherein said one or more ligands are not toxic to said cell.

Title: METHOD FOR MODULATING EXPRESSION OF EXOGENOUS GENES IN MAMMALIAN SYSTEMS, AND PRODUCTS RELATED THERETO

Inventor: EVANS, RONALD M.

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Related patents: 5874534, 6576422, 62245531

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